## STIC Biotechnology Systems Branch

## RAW SEQUENCE LISTING ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number:  $\frac{10/546}{100}$ , Source:  $\frac{10/546}{100}$  Date Processed by STIC:  $\frac{8/30/05}{100}$ 

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.
PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,

TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR CRF SUBMISSION AND PATENTIN SOFTWARE QUESTIONS, PLEASE CONTACT MARK SPENCER, TELEPHONE: 571-272-2510; FAX: 571-273-0221

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE <u>CHECKER VERSION 4.2.2 PROGRAM</u>, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS:

http://www.uspto.gov/web/offices/pac/checker/chkrnote.htm

Applicants submitting genetic sequence information electronically on diskette or CD-Rom should be aware that there is a possibility that the disk/CD-Rom may have been affected by treatment given to all incoming mail. Please consider using alternate methods of submission for the disk/CD-Rom or replacement disk/CD-Rom.

Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

- 1. EFS-Bio (<http://www.uspto.gov/ebc/efs/downloads/documents.htm>, EFS Submission User Manual ePAVE)
- 2. U.S. Postal Service: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450
- 3. Hand Carry, Federal Express, United Parcel Service, or other delivery service (EFFECTIVE 01/14/05): U.S. Patent and Trademark Office. Mail Stop Sequence, Customer Window, Randolph Building. 401 Dulany Street. Alexandria, VA 22314

Revised 01/24/05

ERROR DETECTED	SUGGESTED CORRECTION SERIAL NUMBER: 16/546, 139		
ATTN: NEW RULES CASES: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE			
1Wrapped Nucleics Wrapped Aminos	The number/text at the end of each line "wrapped" down to the next line. This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3; this will prevent "wrapping."		
2Invalid Line Length	The rules require that a line <b>not exceed 72</b> characters in length. This includes white spaces.		
3Misaligned Amino Numbering	The numbering under each 5 <sup>th</sup> amino acid is misaligned. Do not use tab codes between numbers; use space characters, instead.		
4Non-ASCII	The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.		
5Variable Length	Sequence(s) contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.		
6PatentIn 2.0 "bug"	A "bug" in Patentln version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s) Normally, Patentln would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.		
7Skipped Sequences (OLD RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence: (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) (i) SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading) (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) This sequence is intentionally skipped		
	Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to include the skipped sequences.		
8Skipped Sequences (NEW RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence. <210> sequence id number <400> sequence id number 000		
9Use of n's or Xaa's (NEW RULES)	Use of n's and/or Xaa's have been detected in the Sequence Listing.  Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present.  In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.		
Response	Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence		
	Sequence(s) missing the <220> "Feature" and associated numeric identifiers and responses. Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section.  (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32) (Sec. 1.823 of Sequence Rules)		
"bug"	Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.		
13 Misuse of n/Xaa	"n" can only represent a single nucleotide; "Xaa" can only represent a single amino acid		



PCT

DATE: 08/30/2005 RAW SEQUENCE LISTING PATENT APPLICATION: US/10/546,139 TIME: 14:03:41 see item 4 on Even Input Set : A:\PTO.SR.txt Output Set: N:\CRF4\08302005\J546139.raw 3 <110> APPLICANT: Metabolic Explorer 5 <120> TITLE OF INVENTION: Procede de preparation de microorganismes evolues la creation 6 ou la modification de voies metaboliques <130> FILE REFERENCE: D20701/ 345773 C--> 10 <140> CURRENT APPLICATION NUMBER: US/10/546,139 C--> 10 <141> CURRENT FILING DATE: 2005-08-17 10 <150> PRIOR APPLICATION NUMBER: FR 0301924 11 <151> PRIOR FILING DATE: 2003-02-18 Does Not Comply Onected Diskette Neode 13 <150> PRIOR APPLICATION NUMBER: FR 0305768 14 <151> PRIOR FILING DATE: 2003-05-14 16 <150> PRIOR APPLICATION NUMBER: FR 0305769 17 <151> PRIOR FILING DATE: 2003-05-14 19 <150> PRIOR APPLICATION NUMBER: FR 0313054 20 <151> PRIOR FILING DATE: 2003-11-06 22 <160> NUMBER OF SEQ ID NOS: 42 24 <170> SOFTWARE: PatentIn version 3.1 26 <210> SEQ ID NO: 1 27 <211> LENGTH: 100 28 <212> TYPE: DNA ce

Wrothwest Aplanation. Here sour

genetic mo

accatottcg ccagtgccgc gcgggtttct

60 (Se 29 <213> ORGANISM: Artificial Sequence 31 <220> FEATURE: 32 <223> OTHER INFORMATION 34 <400> SEQUENCE: 1 35 taccccegae geaagttetg egeegeetge accatgtteg eeagtgeege gegggtttet 37 ggccagccgc gcgttttcag catatgaata tcctccttag 40 <210> SEQ ID NO: 2 41 <211> LENGTH: 100 42 <212> TYPE: DNA 43 <213> ORGANISM: Artificial Sequence 45 <220> FEATURE: 46 <223> OTHER INFORMATION DmetEF 48 <400> SEQUENCE: 2 60 49 tgacaatatt gaatcacacc ctcggtttcc ctcgcgttgg cctgcgtcgc gagctgaaaa 51 aagcgcaaga aagttattgg tgtaggctgg agctgcttcg 100 54 <210> SEQ ID NO: 3 55 <211> LENGTH: 30 56 <212> TYPE: DNA 57 <213> ORGANISM: Artificial Sequence 59 <220> FEATURE: 60 <223> OTHER INFORMATION: MetER 62 <400> SEQUENCE: 3

66 <210> SEQ ID NO: 4

63 ggtttaagca gtatggtggg aagaagtcgc

30

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

```
67 <211> LENGTH: 30
68 <212> TYPE: DNA
69 <213> ORGANISM: Artificial Sequence
71 <220> FEATURE:
72 <223> OTHER INFORMATION
                            MetEF
74 <400> SEQUENCE: 4
75 cccggggatg aataaacttg ccgccttccc
                                                                          30
78 <210> SEQ ID NO: 5
79 <211> LENGTH: 1161
80 <212> TYPE: DNA
81 <213> ORGANISM: Escherichia coli
83 <400> SEQUENCE: 5
84 atgacgcgta aacaggccac catcgcagtg cgtagcgggt taaatgacga cgaacagtat
                                                                          60
86 ggttgcgttg tcccaccgat ccatctttcc agcacctata actttaccgg atttaatgaa
                                                                         120
88 ccgcgcgcgc atgattactc gcgtcgcggc aacccaacgc gcgatgtggt tcagcgtgcg
                                                                         180
90 ctggcagaac tggaaggtgg tgctggtgca gtacttacta ataccggcat gtccgcgatt
                                                                         240
92 cacctggtaa cgaccgtctt tttgaaacct ggcgatctgc tggttgcgcc gcacgactgc
                                                                         300
94 tacggcggta gctatcgcct gttcgacagt ctggcgaaac gcggttgcta tcgcgtgttg
                                                                         360
96 tttgttgatc aaggcgatga acaggcatta cgggcagcgc tggcagaaaa acccaaactg
                                                                         420
98 gtactggtag aaagcccaag taatccattg ttacgcgtcg tggatattgc gaaaatctgc
                                                                         480
100 catctggcaa gggaagtcgg ggcggtgagc gtggtggata acaccttctt aagcccggca
                                                                          540
102 ttacaaaatc cgctggcatt aggtgccgat ctggtgttgc attcatgcac gaaatatctg
                                                                          600
104 aacggtcact cagacgtagt ggccggcgtg gtgattgcta aagacccgga cgttgtcact
                                                                          660
106 gaactggcct ggtgggcaaa caatattggc gtgacgggg gcgcgtttga cagctatctg
                                                                          720
108 ctgctacgtg ggttgcgaac gctggtgccg cgtatggagc tggcgcagcg caacgcgcag
                                                                          780
110 gcgattgtga aatacctgca aacccagccg ttggtgaaaa aactgtatca cccgtcgttg
                                                                          840
                                                                          900
112 ccggaaaatc aggggcatga aattgccgcg cgccagcaaa aaggctttgg cgcaatgttg
114 agttttgaac tggatggcga tgagcagacg ctgcgtcgtt tcctgggcgg gctgtcgttg
                                                                          960
116 tttacgctgg cggaatcatt agggggagtg gaaagtttaa tctctcacgc cgcaaccatg
                                                                         1020
118 acacatgcag gcatggcacc agaagcgcgt gctgccgccg ggatctccga gacgctgctg
                                                                         1080
120 cgtatctcca ccggtattga agatggcgaa gatttaattg ccgacctgga aaatggcttc
                                                                         1140
122 cgggctgcaa acaaggggta a
                                                                         1161
125 <210> SEQ ID NO: 6
126 <211> LENGTH: 10
127 <212> TYPE: PRT
128 <213> ORGANISM: Escherichia coli
130 <400> SEQUENCE: 6
132 Met Glu Thr Thr His Arg Ala Arg Gly Leu
                                        10
133 1
136 <210> SEQ ID NO: 7
137 <211> LENGTH: 1161
138 <212> TYPE: DNA
139 <213> ORGANISM: Escherichia coli
141 <400> SEQUENCE: 7
142 atgacgcgta aacaggccac catcgcagtg cgtagcgggt taaatgacga cgaacagtat
                                                                           60
                                                                          120
144 ggttgcgttg tcccaccgat ccatctttcc agcacctata actttaccgg atttaatgaa
146 ccgcgcgcgc atgattactc gcgtcgcgqc aacccaacgc gcgatgtggt tcagcgtqcq
                                                                          180
148 ctggcagaac tggaaggtgg tgctggtgca gtacttacta ataccggcat gtccgcgatt
                                                                          240
150 cacctggtaa cgaccgtctt tttgaaacct ggcgatctgc tggttgcgcc gcacgactgc
                                                                          300
```

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

```
152 tacggcggta gctatcgcct gttcgacagt ctggcgaaac gcggttgcta tcgcgtgttg
                                                                           360
154 tttgttgatc aaggcgatga acaggcatta cgggcagcgc tggcagaaaa acccaaactg
                                                                           420
156 gtactggtag aaagcccaag taatccattg ttacgcgtcg tggatattgc gaaaatctgc
                                                                           480
158 catctggcaa gggaagtcgg ggcggtgagc gtggtggata acaccttctt aagcccggca
                                                                           540
160 ttacaaaatc cgctggcatt aggtgccgat ctggtgttgc attcatgcac gaaatatctg
                                                                           600
162 aacggtcact cagacgtagt ggccggcgtg gtgattgcta aagacccgga cgttgtcact
                                                                           660
164 gaactggcct ggtgggcaaa caatattggc gtgacgggcg gcgcgtttga cagctatctg
                                                                           720
166 ctgctacgtg ggttgcgaac gctggtgccg cgtatggagc tggcgcagcg caacgcgcag
                                                                           780
168 gcgattgtga aatacctgca aacccagccg ttggtgaaaa aactgtatca cccgtcgttg
                                                                          840
170 ccggaaaatc aggggcatga aattgccgcg cgccagcaaa aaggctttgg cgcaatgttg
                                                                          900
172 agttttgaac tggatggcga tgagcagacg ctgcgtcgtt tcctgggcgg gctgtcgttg
                                                                          960
174 tttacgctgg cggcatcatt agggggagtg gaaagtttaa tctctcacgc cgcaaccatg
                                                                         1020
176 acacatgcag gcatggcacc agaagcgcgt gctgccgccg ggatctccga gacgctgctg
                                                                         1080
178 cgtatctcca ccggtattga agatggcgaa gatttaattg ccgacctgga aaatggcttc
                                                                         1140
180 cgggctgcaa acaaggggta a
                                                                          1161
183 <210> SEQ ID NO: 8
184 <211> LENGTH: 5
185 <212> TYPE: PRT
186 <213> ORGANISM: Escherichia coli
188 <400> SEQUENCE: 8
190 Met Glu Thr Thr His
191 1
194 <210> SEQ ID NO: 9
195 <211> LENGTH: 30
196 <212> TYPE: DNA
197 <213> ORGANISM: Artificial Sequence
199 <220> FEATURE:
200 <223> OTHER INFORMATION
                             MetJR
202 <400> SEQUENCE: 9
                                                                            30
203 ggtacagaaa ccagcaggct gaggatcagc
206 <210> SEQ ID NO: 10
207 <211> LENGTH: 100
208 <212> TYPE: DNA
209 <213> ORGANISM: Artificial Sequence
211 <220> FEATURE:
212 <223> OTHER INFORMATION: DmetJBF
214 <400> SEQUENCE: 10
215 tatgcagctg acgacettte geceetgeet gegeaateac acteattttt acceettgtt
                                                                            60
                                                                           100
217 tgcagcccgg aagccatttt caggcaccag agtaaacatt
220 <210> SEQ ID NO: 11
221 <211> LENGTH: 30
222 <212> TYPE: DNA
223 <213> ORGANISM: Artificial Sequence
225 <220> FEATURE:
226 <223> OTHER INFORMATION: MetCR
228 <400> SEQUENCE: 11
                                                                            30
229 cgtccgggac gccttgatcc cggacgcaac
232 <210> SEQ ID NO: 12
233 <211> LENGTH: 32
```

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

```
234 <212> TYPE: DNA
235 <213> ORGANISM: Artificial Sequence
237 <220> FEATURE:
                            MetCF
238 <223> OTHER INFORMATION;
240 <400> SEQUENCE: 12
                                                                            32
241 gcgtttacgc agtaaaaaag tcaccagcac gc
244 <210> SEQ ID NO: 13
245 <211> LENGTH: 100
246 <212> TYPE: DNA
247 <213> ORGANISM: Artificial Sequence
249 <220> FEATURE:
250 <223> OTHER INFORMATION: DmetCR
252 <400> SEQUENCE: 13
253 coggogtoca gatoggoaat cagatogtog acatottoca gaccaatatg caggogaato
                                                                            60
255 aaggtcccgc taaaatcgat catatgaata tcctccttag
                                                                           100
258 <210> SEQ ID NO: 14
259 <211> LENGTH: 100
260 <212> TYPE: DNA
261 <213> ORGANISM: Artificial Sequence
263 <220> FEATURE:
264 <223> OTHER INFORMATION: DmetCF
266 <400> SEQUENCE: 14
267 cggacaaaaa gcttgatact caactggtga atgcaggacg cagcaaaaaa tacactctcg
                                                                            60
                                                                           100
269 gcgcggtaaa tagcgtgatt tgtaggctgg agctgcttcg
272 <210> SEQ ID NO: 15
273 <211> LENGTH: 100
274 <212> TYPE: DNA
275 <213> ORGANISM: Artificial Sequence
277 <220> FEATURE:
278 <223> OTHER INFORMATION
                             DcysKR
280 <400> SEQUENCE: 15
281 tgttgcaatt ctttctcagt gaagagatcg gcaaacaatg cggtgcttaa ataacgctca
                                                                            60
283 cccgatgatg gtagaataac catatgaata tcctccttag
                                                                           100
286 <210> SEQ ID NO: 16
287 <211> LENGTH: 100
288 <212> TYPE: DNA
289 <213> ORGANISM: Artificial Sequence
291 <220> FEATURE:
292 <223> OTHER INFORMATION (
                             DcysKF
294 <400> SEQUENCE: 16
295 agtaagattt ttgaagataa ctcgctgact atcggtcaca cgccgctggt tcgcctgaat
                                                                            60
297 cgcatcggta acggacgcat tgtaggctgg agctgcttcg
                                                                           100
300 <210> SEQ ID NO: 17
301 <211> LENGTH: 100
302 <212> TYPE: DNA
303 <213> ORGANISM: Artificial Sequence
305 <220> FEATURE:
306 <223> OTHER INFORMATION; DCysMR
308 <400> SEQUENCE: 17
```

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

309	cccgccccct ggctaaaatg ctcttcccca aacaccccgg tagaaaggta gcgatcgcca	60
	cgatcgcaga tgatcgccac catatgaata tcctccttag	100
	<210> SEQ ID NO: 18	
	<211> LENGTH: 100	
	<212> TYPE: DNA	
	<213> ORGANISM: Artificial Sequence	
	<220> FEATURE:	
	<223> OTHER INFORMATION: DCYSMF	
	<400> SEQUENCE: 18	
	agtacattag aacaaacaat aggcaatacg cctctggtga agttgcagcg aatggggccg	60
	gataacggca gtgaagtgtg tgtaggctgg agctgcttcg	100
	<210> SEQ ID NO: 19	100
	<211> SEQ 1D NO: 19 <211> LENGTH: 30	
	<212> TYPE: DNA	
	<213> ORGANISM: Artificial Sequence	
	<220> FEATURE:	
	<223> OTHER INFORMATION CYSKR	
	<400> SEQUENCE: 19	
	tttttaacag acgcgacgca cgaagagcgc	30
	<210> SEQ ID NO: 20	
	<211> LENGTH: 30	
	<212> TYPE: DNA	
	<213> ORGANISM: Artificial Sequence	
	<220> FEATURE:	
	<223> OTHER INFORMATION (cyskr)	
348 <	<400> SEQUENCE: 20	
349	ggcgcgacgg cgatgtgggt cgattgctat	30
352 <	<210> SEQ ID NO: 21	
353 <	<211> LENGTH: 30	
354 <	<212> TYPE: DNA	
355 <	<213> ORGANISM: Artificial Sequence	
357 <	<220> FEATURE:	
358 <	<223> OTHER INFORMATION, CYSMR	
360 <	<400> SEQUENCE: 21 ( - )	
361	ggggtgacgg tcaggactca ccaatacttc	30
	<210> SEQ ID NO: 22	
	<211> LENGTH: 30	
	<212> TYPE: DNA	
	<213> ORGANISM: Artificial Sequence	
	<220> FEATURE:	
	<223> OTHER INFORMATION CYSMF	
	<400> SEQUENCE: 22	
	gegegeateg etggeegetg ggetacacae	30
	<210> SEQ ID NO: 23	50
	<211> LENGTH: 74	
	COLOS TUDE DUE	
	<212> TIFE: DNA <213> ORGANISM: Artificial Sequence	
	<220> FEATURE: Quotal lun	
	<223> OTHER INFORMATION (Metyr)	
302 <	<pre>&lt;212&gt; TYPE: DNA &lt;213&gt; ORGANISM: Artificial Sequence &lt;220&gt; FEATURE: &lt;223&gt; OTHER INFORMATION MetyR The types of cross shown exist three tips Sequence</pre>	
	the Segrence shown exist three	Chouse

the Sequence Using. Please check subsequent sequences for similar cubrs.

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

## Invalid Line Length:

The rules require that a line not exceed 72 characters in length. This includes spaces.

Seq#:1; Line(s) 5

VERIFICATION SUMMARY

DATE: 08/30/2005

PATENT APPLICATION: US/10/546,139 TIME: 14:03:42

Input Set : A:\PTO.SR.txt

Output Set: N:\CRF4\08302005\J546139.raw

L:10 M:270 C: Current Application Number differs, Replaced Current Application No L:10 M:271 C: Current Filing Date differs, Replaced Current Filing Date